Amendments to the Claims:

This listing of claims will replace all prior versions and listings of claims in the application:

Listing of Claims:

- 1. (Canceled)
- 2. (Currently amended) A method for suppressing the immune system T cell function of an animal, comprising administering to the animal an amount of a hedgehog agonist effective to suppress the immune system T cell function, wherein the hedgehog agonist is a polypeptide which includes a hedgehog amino acid sequence that is at least 90% identical to at least one of SEQ ID Nos. 10-18, or any fragment thereof that binds to a patched polypeptide.
- 3. (Withdrawn) A method for enhancing the immune system of an animal comprising administering to the animal an immunostimulatory amount of a *hedgehog* antagonist.
- 4. (Canceled)
- 5. (Canceled)
- 6. (Currently amended) The method of claim [[4]]2 or 31, wherein the hedgehog amino acid sequence is at least 90 percent identical to at least one of SEQ ID Nos. 10-18 or any fragment thereof that binds to a *patched* polypeptide.
- 7. (Currently amended) The method of claim [[4]]2 or 31, wherein the hedgehog amino acid sequence is encodable by a nucleic acid which hybridizes under stringent conditions of 6.0 x sodium chloride/sodium citrate (SSC) at about 45 °C, followed by a wash of 2.0 x SSC at 50 °C, to at least one of SEQ ID Nos. 1-9.
- 8. (Currently amended) The method of claim [[4]]2 or 31, wherein the hedgehog amino acid sequence is a vertebrate hedgehog polypeptide.
- 9. (Currently amended) The method of claim [[4]]2 or 31, wherein the polypeptide includes at least a 50 amino acid extracellular portion of a vertebrate hedgehog polypeptide.

- 10. (Currently amended) The method of claim [[4]]2 or 31, wherein the polypeptide includes at least an extracellular portion of a vertebrate hedgehog polypeptide corresponding to residues 24-194 of SEQ ID No:15.
- 11. (Currently amended) The method of claim [[4]]2 or 31, wherein the polypeptide is modified with one or more lipophilic moieties.
- 12. (Previously presented) The method of claim 11, wherein the polypeptide is modified with one or more sterol moicties.
- 13. (Previously presented) The method of claim 12, wherein the sterol moiety is cholesterol.
- 14. (Previously presented) The method of claim 11, wherein the polypeptide is modified with one or more fatty acid moietics.
- 15. (Previously presented) The method of claim 14, wherein each fatty acid moiety is independently selected from myristoyl, palmitoyl, stearoyl, and arachidoyl.
- 16. (Previously presented) The method of claim 11, wherein the polypeptide is modified with one or more aromatic hydrocarbons.
- 17. (Previously presented) The method of claim 16, wherein each aromatic hydrocarbon is independently selected from benzene, perylene, phenanthrene, anthracene, naphthalene, pyrene, chrysene, and naphthacene.
- 18. (Previously presented) The method of claim 11, wherein the polypeptide is modified one or more times with a C7 C30 alkyl or cycloalkyl.
- 19. (Withdrawn) The method of claim 1, wherein the *ptc* therapeutic is a small organic molecule.
- 20. (Withdrawn) The method of claim 19, wherein the binding of the ptc therapeutic to patched results in up- or down-regulation of patched and/or gli expression.
- 21. (Previously presented) The method of claim 2 or 31, wherein the hedgehog agonist binds to patched and mimics hedgehog signal transduction by altering the localization, protein-

protein binding, and/or enzymatic activity of an intracellular protein involved in hedgehog signaling.

- 22. (Withdrawn) The method of claim 19, wherein the *ptc* therapeutic is an inhibitor of protein kinase A.
- 23. (Withdrawn) The method of claim 22, wherein the PKA inhibitor is a 5-isoquinolinesulfonamide
- 24. (Withdrawn) The method of claim 22, wherein the PKA inhibitor is represented in the general formula:

wherein,

R1 and R2 cach can independently represent hydrogen, and as valence and stability permit a lower alkyl, a lower alkenyl, a lower alkynyl, a carbonyl (such as a carboxyl, an ester, a formate, or a ketone), a thiocarbonyl (such as a thioester, a thioacetate, or a thioformate), an amino, an acylamino, an amido, a cyano, a nitro, an azido, a sulfate, a sulfonate, a sulfonamido, -(CH₂)_m-R8, -(CH₂)_m-OH, -(CH₂)_m-O-lower alkyl, -(CH₂)_m-O-lower alkyl, -(CH₂)_m-S-lower alkyl, -(CH₂)_m-S-lower alkyl, -(CH₂)_m-S-lower alkyl, -(CH₂)_m-R8, or

R1 and R2 taken together with N form a heterocycle (substituted or unsubstituted);

R3 is absent or represents one or more substitutions to the isoquinoline ring such as a lower alkyl, a lower alkenyl, a lower alkynyl, a carbonyl (such as a carboxyl, an ester, a formate, or a ketone), a thiocarbonyl (such as a thioester, a thioacetate, or a thioformate), an amino, an acylamino, an amido, a cyano, a nitro, an azido, a sulfate, a sulfonate, a

sulfonamido, -(CH₂)_m-R8, -(CH₂)_m-OH, -(CH₂)_m-O-lower alkyl, -(CH₂)_m-O-lower alkenyl, -(CH₂)_n-O-(CH₂)_m-R8, -(CH₂)_m-SH, -(CH₂)_m-S-lower alkyl, -(CH₂)_m-S-lower alkenyl, -(CH₂)_n-S-(CH₂)_m-R8;

R8 represents a substituted or unsubstituted aryl, aralkyl, cycloalkyl, cycloalkenyl, or heterocycle; and

n and m are independently for each occurrence zero or an integer in the range of 1 to 6.

- 25. (Withdrawn) The method of claim 22, wherein the PKA inhibitor is cyclic AMP analog.
- 26. (Withdrawn) The method of claim 22, wherein the PKA inhibitor is selected from the group consisting of N-[2-((p-bromocinnamyl)amino)ethyl]-5-isoquinolinesulfonamide, 1-(5-isoquinoline-sulfonyl)-2-methylpiperazine, KT5720, 8-bromo-cAMP, dibutyryl-cAMP and PKA Heat Stable Inhibitor isoform α.
- 27. (Withdrawn) A therapeutic preparation of a small molecule antagonist of *patched*, which *patched* antagonist is provided in a pharmaceutically acceptable carrier and in an amount sufficient to modulate the immune system of an adult human patient.
- 28. (Withdrawn) A method for modulating T lymphocytes maturation, comprising administering to a patient a gene activation construct which recombines with a genomic *hedgehog* gene of the patient to provide a heterologous transcriptional regulatory sequence operatively linked to a coding sequence of the *hedgehog* gene.
- 29. (Currently amended) The method of claim 2, wherein suppressing the immune system <u>T</u> cell function of an animal comprises inhibiting T lymphocyte maturation in the thymus.
- 30. (Withdrawn) A method of claim 3, wherein enhancing the immune function of an animal comprises stimulating T lymphocyte maturation.
- 31. (Currently amended) A method for suppressing T cell maturation in the thymus, comprising contacting the T cell with an amount of a hedgehog agonist effective to suppress T cell maturation in the thymus, wherein the hedgehog agonist is a polypeptide

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which includes a *hedgehog* amino acid sequence that is at least 90% identical to at least one of SEQ ID Nos. 10-18, or any fragment thereof that binds to a *patched* polypeptide.

- 32. (Canceled)
- 33. (New) The method of claim 2 or 31, wherein the *hedgehog* agonist is an N-terminal fragment of the *hedgehog* polypeptide comprising at least 50 contiguous amino acids.
- 34. (New) The method of claim 33, wherein the *hedgehog* agonist is an N-terminal fragment of the *hedgehog* polypeptide comprising at least 150 contiguous amino acids.
- 35. (New) The method of claim 33, wherein the *hedgehog* agonist includes at least an N-terminal portion of the *hedgehog* polypeptide corresponding to a 19 kDa fragment of an extracellular domain of the *hedgehog* polypeptide.